

NEWS 1	Web Page URLs for STN Seminar Schedule - N. America	
NEWS 2	Apr 08	"Ask CAS" for self-help around the clock
NEWS 3	Apr 09	BEILSTEIN: Reload and Implementation of a New Subject Area
NEWS 4	Apr 09	ZDB will be removed from STN
NEWS 5	Apr 19	US Patent Applications available in IFICDB, IFIPAT, and IFIUDB
NEWS 6	Apr 22	Records from IP.com available in CAPLUS, HCAPLUS, and ZCAPLUS
NEWS 7	Apr 22	BIOSIS Gene Names now available in TOXCENTER
NEWS 8	Apr 22	Federal Research in Progress (FEDRIP) now available
NEWS 9	Jun 03	New e-mail delivery for search results now available
NEWS 10	Jun 10	MEDLINE Reload
NEWS 11	Jun 10	PCTFULL has been reloaded
NEWS 12	Jul 02	FOREGE no longer contains STANDARDS file segment
NEWS 13	Jul 22	USAN to be reloaded July 28, 2002; saved answer sets no longer valid
NEWS 14	Jul 29	Enhanced polymer searching in REGISTRY
NEWS 15	Jul 30	NETFIRST to be removed from STN
NEWS 16	Aug 08	CANCERLIT reload
NEWS 17	Aug 08	PHARMAMarketLetter(PHARMAML) - new on STN
NEWS 18	Aug 08	NTIS has been reloaded and enhanced
NEWS 19	Aug 19	Aquatic Toxicity Information Retrieval (AQUIRE) now available on STN
NEWS 20	Aug 19	IFIPAT, IFICDB, and IFIUDB have been reloaded
NEWS 21	Aug 19	The MEDLINE file segment of TOXCENTER has been reloaded
NEWS 22	Aug 26	Sequence searching in REGISTRY enhanced
NEWS 23	Sep 03	JAPIO has been reloaded and enhanced
NEWS 24	Sep 16	Experimental properties added to the REGISTRY file
NEWS 25	Sep 16	Indexing added to some pre-1967 records in CA/CAPLUS
NEWS 26	Sep 16	CA Section Thesaurus available in CAPLUS and CA
NEWS 27	Oct 01	CASREACT Enriched with Reactions from 1907 to 1985

NEWS EXPRESS February 1 CURRENT WINDOWS VERSION IS V6.0d,
CURRENT MACINTOSH VERSION IS V6.0a(ENG) AND V6.0Ja(JP),
AND CURRENT DISCOVER FILE IS DATED 05 FEBRUARY 2002

NEWS HOURS STN Operating Hours Plus Help Desk Availability

NEWS INTER General Internet Information

NEWS LOGIN Welcome Banner and News Items

NEWS LOGIN **NEWS PHONE** **WELCOME** **BANNER** and **NEWS** **ITEMS**
Direct Dial and Telecommunica

NEWS PHONE CAS World Wide Web Site (general information)

NEWS WWW.CBS.WORLD.WIDE.WEB.BY.CBS (GENERAL INFORMATION)

Enter NEWS followed by the item number or name to see news on that specific topic.

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FILE 'HOME' ENTERED AT 20:08:06 ON 03 OCT 2002

=> file reg
COST IN U.S. DOLLARS
SINCE FILE
ENTRY
TOTAL
SESSION
0.21
0.21
FULL ESTIMATED COST

FILE 'REGISTRY' ENTERED AT 20:08:18 ON 03 OCT 2002 :
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Property values tagged with IC are from the ZIC/VINITI data file provided by InfoChem.

STRUCTURE FILE UPDATES: 2 OCT 2002 HIGHEST RN 458522-67-3
 DICTIONARY FILE UPDATES: 2 OCT 2002 HIGHEST RN 458522-67-3

TSCA INFORMATION NOW CURRENT THROUGH MAY 20, 2002

Please note that search-term pricing does apply when conducting SmartSELECT searches.

Crossover limits have been increased. See HELP CROSSOVER for details.

Experimental and calculated property data are now available. See HELP PROPERTIES for more information. See STNote 27, Searching Properties in the CAS Registry File, for complete details:
<http://www.cas.org/ONLINE/STN/STNOTES/stnotes27.pdf>

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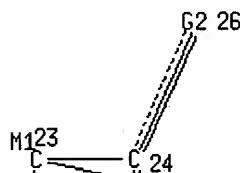
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L1 HAS NO ANSWERS

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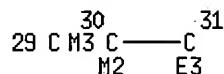
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Page 1-A

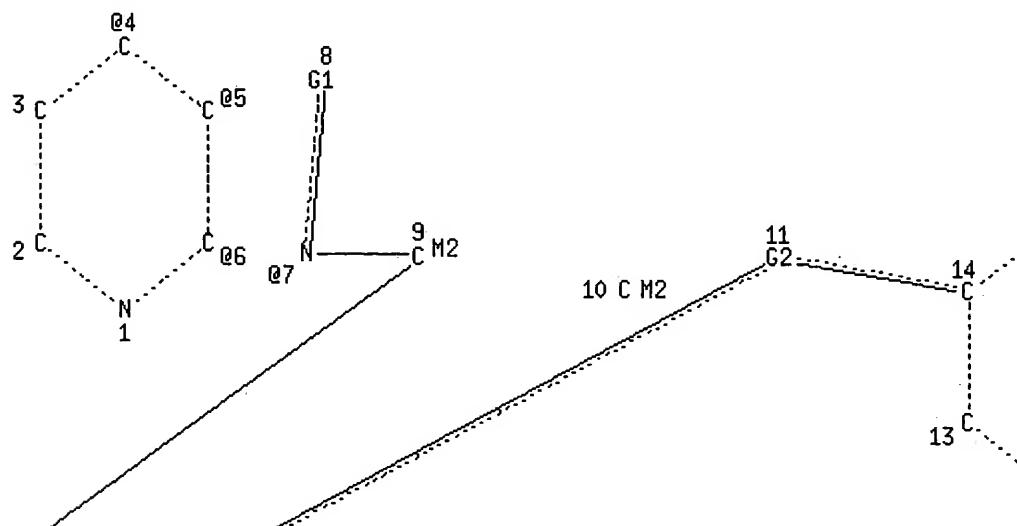


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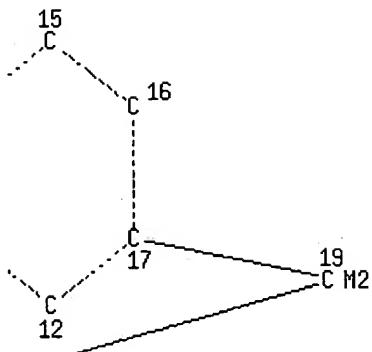
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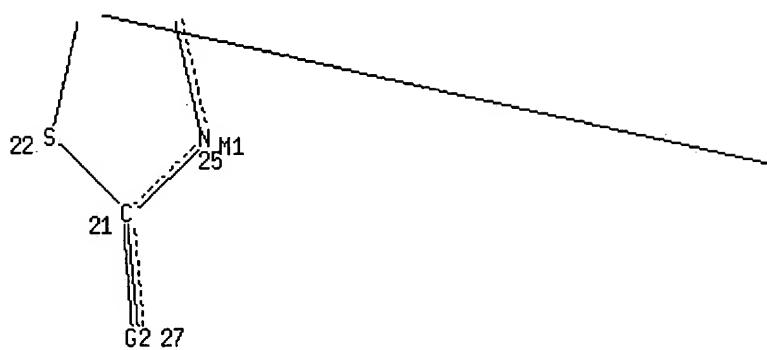
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Page 1-E

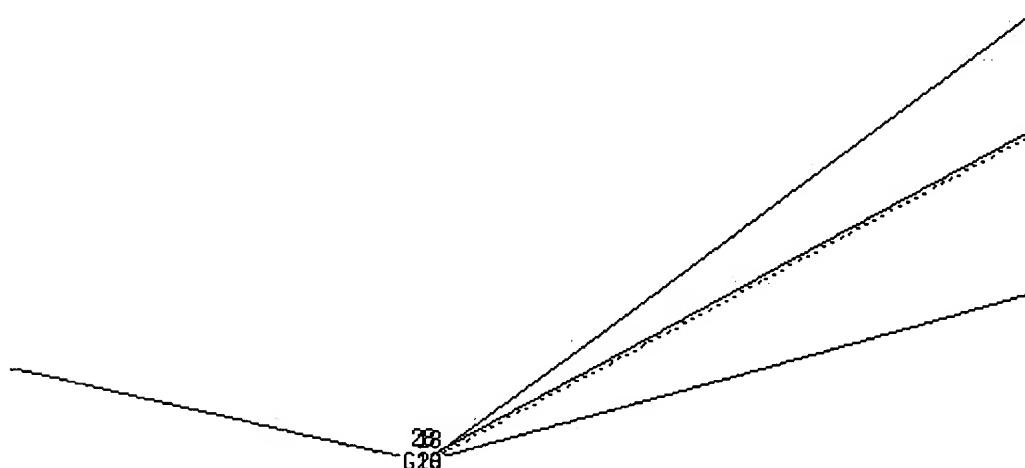


Page 1-F



Page 2-B

Page 2-C



Page 2-D

Page 2-E

Page 2-F

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VAR G2=32/33

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REP G20=(0-1) 10-9 10-11

VPA 7-4/5/6 S

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HCOUNT IS M2 AT 19

HCOUNT IS M2 AT 20

HCOUNT IS M1 AT 23

HCOUNT IS M1 AT 25

HCOUNT IS M3 AT 29

HCOUNT IS M2 AT 30

HCOUNT IS E3 AT 31

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DEFAULT ECLEVEL IS LIMITED

GRAPH ATTRIBUTES:

RSPEC I
NUMBER OF NODES IS 33

STEREO ATTRIBUTES: NONE

=> **s 11**
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SAMPLE SCREEN SEARCH COMPLETED - 10 TO ITERATE

100.0% PROCESSED 10 ITERATIONS 2 ANSWERS
SEARCH TIME: 00.00.02

FULL FILE PROJECTIONS: ONLINE **COMPLETE**
BATCH **COMPLETE**
PROJECTED ITERATIONS: 11 TO 389
PROJECTED ANSWERS: 2 TO 124

L2 2 SEA SSS SAM L1

=> **s 11 full**
THE ESTIMATED SEARCH COST FOR FILE 'REGISTRY' IS 139.90 U.S. DOLLARS
DO YOU WANT TO CONTINUE WITH THIS REQUEST? (Y)/N or END:y
FULL SEARCH INITIATED 20:14:18 FILE 'REGISTRY'
FULL SCREEN SEARCH COMPLETED - 310 TO ITERATE

100.0% PROCESSED 310 ITERATIONS 48 ANSWERS
SEARCH TIME: 00.00.06

L3 48 SEA SSS FUL L1

=> **file hcaplus**
COST IN U.S. DOLLARS SINCE FILE TOTAL
FULL ESTIMATED COST ENTRY SESSION
143.70 143.91

FILE 'HCAPLUS' ENTERED AT 20:14:30 ON 03 OCT 2002
USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT.
PLEASE SEE "HELP USAGETERMS" FOR DETAILS.
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FILE COVERS 1907 - 3 Oct 2002 VOL 137 ISS 14
FILE LAST UPDATED: 2 Oct 2002 (20021002/ED)

This file contains CAS Registry Numbers for easy and accurate substance identification.

CAS roles have been modified effective December 16, 2001. Please check your SDI profiles to see if they need to be revised. For information on CAS roles, enter HELP ROLES at an arrow prompt or use the CAS Roles thesaurus (/RL field) in this file.

=> **s 13**
L4 490 L3

=> s 13/thu
 490 L3
 466819 THU/RL
 L5 324 L3/THU
 (L3 (L) THU/RL)

=> s 15 and method
 2340863 METHOD
 982149 METHODS
 3048039 METHOD
 (METHOD OR METHODS)
 L6 92 L5 AND METHOD

=> s 16 and diab?
 94540 DIAB?
 L7 56 L6 AND DIAB?

=> s 17 and mellit?
 53014 MELLIT?
 L8 44 L7 AND MELLIT?

=> s 18 and pd < may 20 1999
 19621278 PD < MAY 20 1999
 (PD<19990520)
 L9 17 L8 AND PD < MAY 20 1999

=> d 19, ibib abs fhitstr, 1-17

L9 ANSWER 1 OF 17 HCAPLUS COPYRIGHT 2002 ACS

Full Citing
 Text References

ACCESSION NUMBER: 2000:362595 HCAPLUS
 DOCUMENT NUMBER: 133:13403
 TITLE: Adipocyte containing ob gene promoter for screening
 modulators useful in treatment of anorexia, obesity,
 and other diseases
 INVENTOR(S): Briggs, Michael R.; Auwerx, Johan; De Vos, Piet;
 Staels, Bart; Croston, Glenn E.; Miller, Stephen G.
 PATENT ASSIGNEE(S): Ligand Pharmaceuticals Inc., USA
 SOURCE: U.S., 64 pp., Cont.-in-part of U.S. Ser. No. 558,588,
 abandoned.
 CODEN: USXXAM
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 2
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
US 6068976	A	20000530	US 1996-618100	19960319
CA 2215387	AA	19960926	CA 1996-2215387	19960319
<u>PRIORITY APPLN. INFO.:</u>				
US 1995-408584 B2 19950320				
US 1995-418096 B2 19950405				
US 1995-510584 B2 19950802				
US 1995-558588 B2 19951030				
US 1995-7390P P 19951121				
US 1995-7721P P 19951130				
US 1995-8601P P 19951214				

AB This invention relates to the isolation and cloning of the promoter and other control regions of a human ob gene. It provides a **method** for identifying and screening for agents useful for the treatment of diseases and pathol. conditions affected by the level of expression of an ob gene. These agents interact directly or indirectly with the promoter or other

control regions of the ob gene. A PPAR γ agonist, BRL49653, has been identified to be useful in treating anorexia, cachexia, and other diseases characterized by insufficient food intake or body wt. loss. Modulators of ob gene expression may be used to treat other diseases such as obesity, **diabetes**, hypertension, cardiovascular diseases and infertility.

IT 122320-73-4, BRL49653

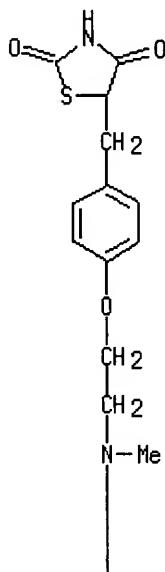
RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(PPAR γ agonist; adipocyte contg. ob gene promoter for screening modulators useful in treatment of anorexia, obesity, and other diseases)

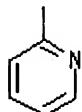
RN 122320-73-4 HCAPLUS

CN 2,4-Thiazolidinedione, 5-[[4-[2-(methyl-2-pyridinylamino)ethoxy]phenyl]methoxy]-(9CI) (CA INDEX NAME)

PAGE 1-A



PAGE 2-A



REFERENCE COUNT:

9

THERE ARE 9 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L9 ANSWER 2 OF 17 HCAPLUS COPYRIGHT 2002 ACS

Full Text Citing References

ACCESSION NUMBER:

1999:316557 HCAPLUS

DOCUMENT NUMBER:

130:332912

TITLE:

Activators of the nuclear orphan receptor peroxisome proliferator-activated receptor gamma for treatment of **diabetes** and cardiovascular disorders

INVENTOR(S):

Kliewer, Steven Anthony; Lehmann, Jurgen M.; Willson, Timothy M.

PATENT ASSIGNEE(S):

Glaxo Wellcome Inc., USA

SOURCE:

U.S., 9 pp., Cont. of U.S. Ser. No. 804,310, abandoned.

CODEN: USXXAM
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 1
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
<u>US 5902726</u>	A	19990511	<u>US 1998-28988</u>	19980225
<u>US 5994554</u>	A	19991130	<u>US 1998-207936</u>	19981209
<u>PRIORITY APPLN. INFO.:</u>			<u>US 1994-363482</u>	19941223
			<u>US 1995-386394</u>	19950210
			<u>US 1997-804310</u>	19970221
			<u>US 1998-28988</u>	19980225

OTHER SOURCE(S): MARPAT 130:332912

AB The present invention provides activator compds., including agonists, to the peroxisome proliferator-activated receptor gamma. Particular PPAR γ activators are set forth, as are a pharmaceutical compn. for treating **diabetes**, non-insulin-dependent **diabetes mellitus**, cardiovascular disorders, and **methods** for such treatment. Also claimed is a **method** of identifying activator compds.

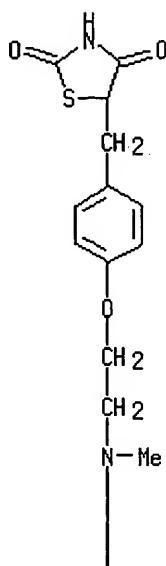
IT 173792-21-7

RL: ARG (Analytical reagent use); THU (Therapeutic use); BSU (Biological study, unclassified); THU (Therapeutic use); ANST (Analytical study); BIOL (Biological study); PROC (Process); USES (Uses) (for identifying PPAR γ -interacting compds. useful as drugs; activators of peroxisome proliferator-activated receptor gamma for treatment of **diabetes** and cardiovascular disorders)

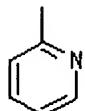
RN 173792-21-7 HCPLUS

CN 2,4-Thiazolidinedione, 5-[[4-[2-(methyl-2-pyridinylamino)ethoxy]phenyl]methyl]-, labeled with tritium (9CI) (CA INDEX NAME)

PAGE 1-A



PAGE 2-A



REFERENCE COUNT: 23 THERE ARE 23 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L9 ANSWER 3 OF 17 HCAPLUS COPYRIGHT 2002 ACS

Full Citing
Text References

ACCESSION NUMBER: 1999:167677 HCAPLUS
 DOCUMENT NUMBER: 131:124868
 TITLE: Systemic exposure to rosiglitazone is unaltered by food
 AUTHOR(S): Freed, M. I.; Allen, A.; Jorkasky, D. K.; DiCicco, R. A.
 CORPORATE SOURCE: SmithKline Beecham Clinical Pharmacology Unit, Presbyterian Medical Center of the University of Pennsylvania Health System, 51 North 39th Street, Philadelphia, PA, 19104, USA
 SOURCE: European Journal of Clinical Pharmacology (1999), 55(1), 53-56
 CODEN: EJCPAS; ISSN: 0031-6970

PUBLISHER: Springer-Verlag
 DOCUMENT TYPE: Journal
 LANGUAGE: English

AB Objective: To evaluate the effect of food on the bioavailability and pharmacokinetics of the insulin sensitizer rosiglitazone. Methods: In a randomized, open-label, period-balanced, single-dose, crossover study, rosiglitazone 2 mg was administered to 12 healthy male volunteers either in the fasting state or following a std. high-fat breakfast. The primary end points of the study were AUC0-inf and Cmax. Results: Single oral doses of rosiglitazone were safe and well tolerated. Overall exposure to rosiglitazone was unaffected by food. The geometric mean ratio of AUC(0-inf) in the fed:fasted regimens was 0.94 (95% CI: 0.82, 1.06); t1/2 was unaffected. Absorption of rosiglitazone in the fed state was more gradual and sustained than in the fasted state. Cmax was reduced by approx. 20% (point est. 0.80; 95% CI 0.65 to 0.97) and tmax was modestly delayed in the fed state. Conclusion: These data support dosing guidelines that will permit the administration of rosiglitazone without regard to meals for treatment of type 2 diabetes mellitus.

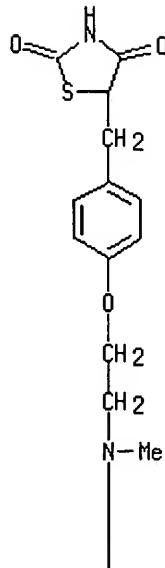
IT 122320-73-4, Rosiglitazone

RL: BPR (Biological process); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); PROC (Process); USES (Uses)
 (bioavailability of antidiabetic rosiglitazone is unaltered by food intake in humans)

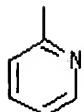
RN 122320-73-4 HCAPLUS

CN 2,4-Thiazolidinedione, 5-[[4-[2-(methyl-2-pyridinylamino)ethoxy]phenyl]methyl]- (9CI) (CA INDEX NAME)

PAGE 1-A



PAGE 2-A



REFERENCE COUNT:

21 THERE ARE 21 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L9 ANSWER 4 OF 17 HCPLUS COPYRIGHT 2002 ACS

Full Text	Citing References
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ACCESSION NUMBER:

1999:81575 HCPLUS

DOCUMENT NUMBER:

130:134189

TITLE:

Treatment of **diabetes** with a thiazolidinedione, an insulin secretagogue, and an α -glucosidase inhibitor

INVENTOR(S):

Buckingham, Robin Edwin; Smith, Stephen Alistair

PATENT ASSIGNEE(S):

Smithkline Beecham PLC, UK

SOURCE:

PCT Int. Appl., 20 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
<u>WO 9903478</u>	<u>A1</u>	<u>19990128</u>	<u>WO 1998-GB2112</u>	<u>19980716</u>
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<u>AU 9884490</u>	<u>A1</u>	<u>19990210</u>	<u>AU 1998-84490</u>	<u>19980716</u>
<u>EP 1001784</u>	<u>A1</u>	<u>20000524</u>	<u>EP 1998-935129</u>	<u>19980716</u>

R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
IE, SI, FI, RO

BR 9810292	A	20000919	BR 1998-10292	19980716
JP 2001510160	T2	20010731	JP 2000-502777	19980716
ZA 9806364	A	20000117	ZA 1998-6364	19980717
NO 2000000230	A	20000117	NO 2000-230	20000117
US 2002052324	A1	20020502	US 2001-989572	20011120

PRIORITY APPLN. INFO.:

GB 1997-15298	A	19970718
WO 1998-GB2112	W	19980716
US 1999-445908	A1	19991215

AB A **method** and compn. are disclosed for the treatment of **diabetes mellitus** and conditions assocd. with **diabetes mellitus** in a mammal. The **method** comprises administering an effective nontoxic and pharmaceutically acceptable amt. of an insulin sensitizer, an insulin secretagogue and an α -glucosidase inhibitor antihyperglycemic agent to a mammal in need thereof.

IT 122320-73-4

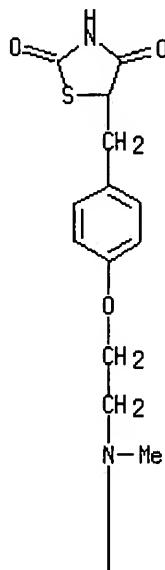
RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(thiazolidinedione, insulin secretagogue, and α -glucosidase inhibitor for **diabetes** treatment)

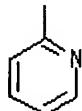
RN 122320-73-4 HCAPLUS

CN 2,4-Thiazolidinedione, 5-[[4-[2-(methyl-2-pyridinylamino)ethoxy]phenyl]met hyl]- (9CI) (CA INDEX NAME)

PAGE 1-A



PAGE 2-A



REFERENCE COUNT:

7

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L9 ANSWER 5 OF 17 HCAPLUS COPYRIGHT 2002 ACS

Full Text	Citing References
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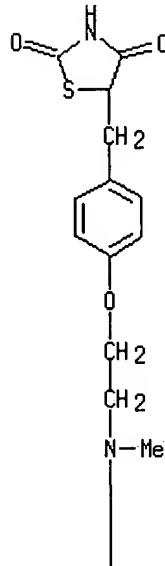
ACCESSION NUMBER: 1999:81574 HCPLUS
 DOCUMENT NUMBER: 130:134188
 TITLE: Treatment of **diabetes** with a thiazolidinedione, an insulin secretagogue, and a biguanide
 INVENTOR(S): Buckingham, Robin Edwin; Smith, Stephen Alistair
 PATENT ASSIGNEE(S): Smithkline Beecham PLC, UK
 SOURCE: PCT Int. Appl., 19 pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 1
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
<u>WO 9903477</u>	<u>A1</u>	<u>19990128</u>	<u>WO 1998-GB2110</u>	<u>19980716</u>
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RW: GH, GM, KE, LS, MW, SD, SZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG				
<u>AU 9884488</u>	<u>A1</u>	<u>19990210</u>	<u>AU 1998-84488</u>	<u>19980716</u>
<u>EP 1001783</u>	<u>A1</u>	<u>20000524</u>	<u>EP 1998-935127</u>	<u>19980716</u>
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, FI, RO				
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<u>JP 2001510159</u>	<u>T2</u>	<u>20010731</u>	<u>JP 2000-502776</u>	<u>19980716</u>
<u>ZA 9806363</u>	<u>A</u>	<u>20000117</u>	<u>ZA 1998-6363</u>	<u>19980717</u>
<u>NO 2000000228</u>	<u>A</u>	<u>20000117</u>	<u>NO 2000-228</u>	<u>20000117</u>
<u>US 2002016287</u>	<u>A1</u>	<u>20020207</u>	<u>US 2001-939470</u>	<u>20010824</u>
<u>PRIORITY APPLN. INFO.:</u>			<u>GB 1997-15295</u>	<u>A</u> <u>19970718</u>
			<u>WO 1998-GB2110</u>	<u>W</u> <u>19980716</u>
			<u>US 1999-446039</u>	<u>A1</u> <u>19991215</u>

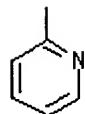
AB A **method** and compn. are disclosed for the treatment of **diabetes mellitus** and conditions assocd. with **diabetes mellitus** in a mammal. The **method** comprises administering an effective nontoxic and pharmaceutically acceptable amt. of an insulin sensitizer, an insulin secretagogue and a biguanide antihyperglycemic agent to a mammal in need thereof.

IT 122320-73-4
 RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
 (thiazolidinedione, insulin secretagogue, and biguanide for **diabetes** treatment)
 RN 122320-73-4 HCPLUS
 CN 2,4-Thiazolidinedione, 5-[[4-[2-(methyl-2-pyridinylamino)ethoxy]phenyl]methyl]- (9CI) (CA INDEX NAME)

PAGE 1-A



PAGE 2-A



REFERENCE COUNT: 7 THERE ARE 7 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L9 ANSWER 6 OF 17 HCAPLUS COPYRIGHT 2002 ACS

Full Citing
 Text References

ACCESSION NUMBER: 1999:81573 HCAPLUS
 DOCUMENT NUMBER: 130:134187
 TITLE: Treatment of **diabetes** with insulin sensitizer thiazolidinedione and insulin secretagogue sulfonylurea
 INVENTOR(S): Buckingham, Robin Edwin; Smith, Stephen Alistair
 PATENT ASSIGNEE(S): Smithkline Beecham PLC, UK
 SOURCE: PCT Int. Appl., 19 pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 1
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
<u>WO 9903476</u>	A1	19990128	<u>WO 1998-GB2109</u>	19980716
W: AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE, DK, EE, ES, FI, GB, GE, GH, GM, HR, HU, ID, IL, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM				
RW: GH, GM, KE, LS, MW, SD, SZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG				
<u>AU 9884487</u>	A1	19990210	<u>AU 1998-84487</u>	19980716
<u>AU 743269</u>	B2	20020124		

<u>EP 998291</u>	A1	20000510	<u>EP 1998-935126</u>	19980716
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, FI, RO				
<u>BR 9810904</u>	A	20000926	<u>BR 1998-10904</u>	19980716
<u>JP 2001510158</u>	T2	20010731	<u>JP 2000-502775</u>	19980716
<u>ZA 9806365</u>	A	20000117	<u>ZA 1998-6365</u>	19980717
<u>NO 2000000229</u>	A	20000117	<u>NO 2000-229</u>	20000117
<u>US 2002045649</u>	A1	20020418	<u>US 2001-975883</u>	20011012
<u>PRIORITY APPLN. INFO.:</u>				
GB <u>1997-15306</u> A 19970718				
WO <u>1998-GB2109</u> W 19980716				
US <u>1999-445907</u> A1 19991215				

AB A **method** for the treatment of **diabetes mellitus** and conditions assoacd. with **diabetes mellitus** in a mammal, which **method** comprises administering an effective non-toxic and pharmaceutically acceptable amt. of an insulin sensitizer and a sub-maximal amt. of an insulin secretagogue, to a mammal in need thereof; and a pharmaceutical compn. for use in such **method** are disclosed. The insulin secretagogue is esp. sulfonylurea. The insulin sensitizer is esp. 5-[4-[2-(N-methyl-N-(2-pyridyl)amino)ethoxy]benzyl]thiazolidine-2,4-dione (I). Tablet formulations contg. I maleate are given.

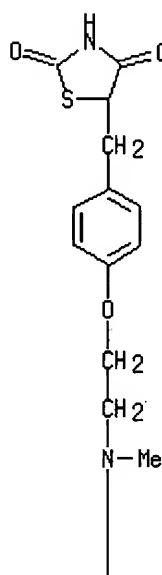
IT 122320-73-4

RL: THU (Therapeutic use); BSU (Biological study, unclassified);
THU (Therapeutic use); BIOL (Biological study); PROC (Process);
USES (Uses)
(as insulin sensitizer; treatment of **diabetes** with insulin sensitizer thiazolidinedione and insulin secretagogue sulfonylurea)

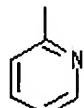
RN 122320-73-4 HCPLUS

CN 2,4-Thiazolidinedione, 5-[(4-[2-(methyl-2-pyridinylamino)ethoxy]phenyl)methyl]- (9CI) (CA INDEX NAME)

PAGE 1-A



PAGE 2-A



REFERENCE COUNT:

8

THERE ARE 8 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L9 ANSWER 7 OF 17 HCPLUS COPYRIGHT 2002 ACS

Full Text	Citing References
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ACCESSION NUMBER: 1999:9712 HCPLUS
 DOCUMENT NUMBER: 130:61091
 TITLE: Treatment of **diabetes** with thiazolidinedione and sulfonylurea
 INVENTOR(S): Smith, Stephen Alistair
 PATENT ASSIGNEE(S): Smithkline Beecham Plc, UK
 SOURCE: PCT Int. Appl., 20 pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 1
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
<u>WO 9857649</u>	<u>A1</u>	<u>19981223</u>	<u>WO 1998-EP3688</u>	<u>19980615</u>
W: AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE, DK, EE, ES, FI, GB, GE, GH, GM, GW, HU, ID, IL, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM				
RW: GH, GM, KE, LS, MW, SD, SZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, ML, MR, NE, SN, TD, TG				
<u>AU 9885392</u>	<u>A1</u>	<u>19990104</u>	<u>AU 1998-85392</u>	<u>19980615</u>
<u>EP 999845</u>	<u>A1</u>	<u>20000517</u>	<u>EP 1998-936363</u>	<u>19980615</u>
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, FI, RO				
<u>BR 9810142</u>	<u>A</u>	<u>20000808</u>	<u>BR 1998-10142</u>	<u>19980615</u>
<u>JP 2001523270</u>	<u>T2</u>	<u>20011120</u>	<u>JP 1999-503754</u>	<u>19980615</u>
<u>ZA 9805236</u>	<u>A</u>	<u>20000217</u>	<u>ZA 1998-5236</u>	<u>19980617</u>
<u>NO 9906264</u>	<u>A</u>	<u>20000217</u>	<u>NO 1999-6264</u>	<u>19991217</u>
<u>US 2001049380</u>	<u>A1</u>	<u>20011206</u>	<u>US 2001-848511</u>	<u>20010502</u>
<u>PRIORITY APPLN. INFO.:</u>			<u>GB 1997-12854</u>	<u>A</u> <u>19970618</u>
			<u>GB 1998-6710</u>	<u>A</u> <u>19980327</u>
			<u>WO 1998-EP3688</u>	<u>W</u> <u>19980615</u>
			<u>US 1999-445859</u>	<u>B1</u> <u>19991215</u>

AB A **method** for the treatment of **diabetes mellitus** and conditions assocd. with **diabetes mellitus** in a mammal, which **method** comprises administering an effective nontoxic and pharmaceutically acceptable amt. of an insulin sensitizer and an insulin secretagogue, to a mammal in need thereof.

IT 155141-29-0, Rosiglitazone maleate

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
 (treatment of **diabetes** with thiazolidinedione and sulfonylurea)

RN 155141-29-0 HCPLUS

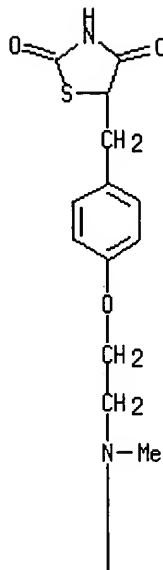
CN 2,4-Thiazolidinedione, 5-[[4-[2-(methyl-2-pyridinylamino)ethoxy]phenyl]methyl]-, (2Z)-2-butenedioate (1:1) (9CI) (CA INDEX NAME)

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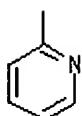
CRN 122320-73-4

CMF C18 H19 N3 O3 S

PAGE 1-A



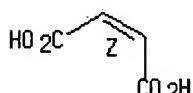
PAGE 2-A



CM 2

CRN 110-16-7
CMF C4 H4 O4

Double bond geometry as shown.



REFERENCE COUNT: 8 THERE ARE 8 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L9 ANSWER 8 OF 17 HCAPLUS COPYRIGHT 2002 ACS

Full	<input checked="" type="checkbox"/> Citing
Text	<input type="checkbox"/> References

ACCESSION NUMBER: 1999:9699 HCAPLUS
 DOCUMENT NUMBER: 130:61090
 TITLE: Treatment of **diabetes** with rosiglitazone and insulin
 INVENTOR(S): Smith, Stephen Alistair
 PATENT ASSIGNEE(S): Smithkline Beecham P.L.C., UK
 SOURCE: PCT Int. Appl., 17 pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 1
PATENT INFORMATION:

PATENT NO. KIND DATE

APPLICATION NO. DATE

<u>WO 9857636</u>	<u>A1 19981223</u>	<u>WO 1998-EP3692</u>	<u>19980615</u>
W: AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE, DK, EE, FI, GB, GE, GH, GM, GW, HU, ID, IL, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM			
RW: GH, GM, KE, LS, MW, SD, SZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, ML, MR, NE, SN, TD, TG			

<u>AU 9882163</u>	<u>A1 19990104</u>	<u>AU 1998-82163</u>	<u>19980615</u>
<u>EP 999837</u>	<u>A1 20000517</u>	<u>EP 1998-932169</u>	<u>19980615</u>
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, FI, RO			
<u>BR 9810444</u>	<u>A 20000905</u>	<u>BR 1998-10444</u>	<u>19980615</u>
<u>JP 2002504138</u>	<u>T2 20020205</u>	<u>JP 1999-503757</u>	<u>19980615</u>
<u>ZA 9805237</u>	<u>A 20000217</u>	<u>ZA 1998-5237</u>	<u>19980617</u>
<u>NO 9906265</u>	<u>A 19991217</u>	<u>NO 1999-6265</u>	<u>19991217</u>
<u>US 2002028768</u>	<u>A1 20020307</u>	<u>US 2001-928326</u>	<u>20010813</u>

PRIORITY APPLN. INFO.:

<u>GB 1997-12866</u>	<u>A 19970618</u>
<u>WO 1998-EP3692</u>	<u>W 19980615</u>
<u>US 1999-445858</u>	<u>B1 19991215</u>

AB A **method** for the treatment of **diabetes mellitus** and conditions assocd. with **diabetes mellitus** in a mammal, which **method** comprises administering an effective nontoxic and pharmaceutically acceptable amt. of insulin sensitizer rosiglitazone and insulin to a mammal in need thereof.

IT 155141-29-0, Rosiglitazone maleate

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(treatment of **diabetes mellitus** with rosiglitazone and insulin)

RN 155141-29-0 HCPLUS

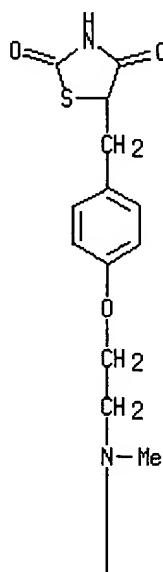
CN 2,4-Thiazolidinedione, 5-[[4-[2-(methyl-2-pyridinylamino)ethoxy]phenyl]methyl]-, (2Z)-2-butenedioate (1:1) (9CI) (CA INDEX NAME)

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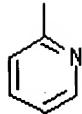
CRN 122320-73-4

CMF C18 H19 N3 O3 S

PAGE 1-A



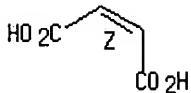
PAGE 2-A



CM 2

CRN 110-16-7
CMF C4 H4 O4

Double bond geometry as shown.



REFERENCE COUNT: 7 THERE ARE 7 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L9 ANSWER 9 OF 17 HCAPLUS COPYRIGHT 2002 ACS

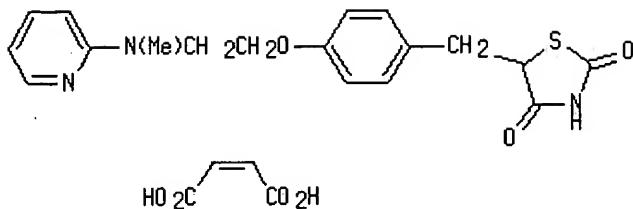
Full Text	Citing References
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ACCESSION NUMBER: 1999:9698 HCAPLUS
 DOCUMENT NUMBER: 130:76189
 TITLE: Treatment of **diabetes** with thiazolidinedione and alpha-glucosidase inhibitor
 INVENTOR(S): Smith, Stephen Alistair
 PATENT ASSIGNEE(S): Smithkline Beecham Plc, UK
 SOURCE: PCT Int. Appl., 19 pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 1
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
<u>WO 9857635</u>	<u>A1</u>	<u>19981223</u>	<u>WO 1998-EP3691</u>	<u>19980615</u>
W: AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE, DK, EE, ES, FI, GB, GE, GH, GM, GW, HU, ID, IL, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM				
RW: GH, GM, KE, LS, MW, SD, SZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, ML, MR, NE, SN, TD, TG				
<u>AU 9887999</u>	<u>A1</u>	<u>19990104</u>	<u>AU 1998-87999</u>	<u>19980615</u>
<u>EP 975343</u>	<u>A1</u>	<u>20000202</u>	<u>EP 1998-939513</u>	<u>19980615</u>
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, FI, RO				
<u>BR 9810186</u>	<u>A</u>	<u>20000808</u>	<u>BR 1998-10186</u>	<u>19980615</u>
<u>JP 2001523271</u>	<u>T2</u>	<u>20011120</u>	<u>JP 1999-503756</u>	<u>19980615</u>
<u>ZA 9805235</u>	<u>A</u>	<u>20000217</u>	<u>ZA 1998-5235</u>	<u>19980617</u>
<u>NO 9906270</u>	<u>A</u>	<u>19991217</u>	<u>NO 1999-6270</u>	<u>19991217</u>
<u>US 2001034356</u>	<u>A1</u>	<u>20011025</u>	<u>US 2001-863136</u>	<u>20010523</u>
<u>US 2002123514</u>	<u>A1</u>	<u>20020905</u>	<u>US 2002-91008</u>	<u>20020305</u>
<u>PRIORITY APPLN. INFO.:</u>				
			<u>GB 1997-12865</u>	<u>A</u> <u>19970618</u>
			<u>GB 1998-6708</u>	<u>A</u> <u>19980327</u>
			<u>WO 1998-EP3691</u>	<u>W</u> <u>19980615</u>

US 1999-445951 B1 19991215
 US 2001-863136 B1 20010523

GI



AB A **method** for the treatment of **diabetes mellitus** and conditions assocd. with **diabetes mellitus** in a mammal, which **method** comprises administering an effective non-toxic and pharmaceutically acceptable amt. of an insulin sensitizer (I) and an α -glucosidase inhibitor antihyperglycemic agent. The effects of α -glucosidase inhibitor acarbose on the pharmacokinetics of I in healthy humans are described along with pharmaceutical formulations (concns. and tablets) contg. I.

IT **155141-29-0**

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(treatment of **diabetes mellitus** and conditions assocd. with **diabetes** with thiazolidinedione deriv. and α -glucosidase inhibitors)

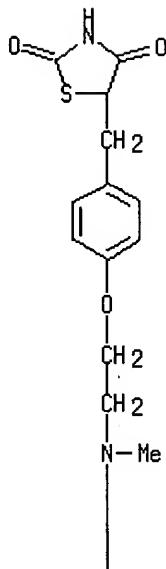
RN **155141-29-0** HCAPLUS

CN 2,4-Thiazolidinedione, 5-[[4-[(2Z)-2-butenedioate]phenoxy]phenyl]methyl-, (2Z)-2-butenedioate (1:1) (9CI) (CA INDEX NAME)

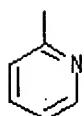
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CRN **122320-73-4**
 CMF C18 H19 N3 O3 S

PAGE 1-A



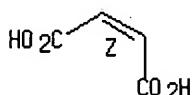
PAGE 2-A



CM 2

CRN 110-16-7
CMF C4 H4 O4

Double bond geometry as shown.



REFERENCE COUNT: 7 THERE ARE 7 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L9 ANSWER 10 OF 17 HCAPLUS COPYRIGHT 2002 ACS

Full	Citing
Text	References

ACCESSION NUMBER: 1999:9697 HCAPLUS
 DOCUMENT NUMBER: 130:61089
 TITLE: Treatment of **diabetes** with thiazolidinedione and metformin
 INVENTOR(S): Smith, Stephen Alistair
 PATENT ASSIGNEE(S): Smithkline Beecham Plc, UK
 SOURCE: PCT Int. Appl., 20 pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 1
PATENT INFORMATION:

PATENT NO. KIND DATE APPLICATION NO. DATE

<u>WO 9857634</u>	<u>A1</u>	<u>19981223</u>	<u>WO 1998-EP3690</u>	<u>19980615</u>
W: AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE, DK, EE, FI, GB, GE, GH, GM, GW, HU, ID, IL, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM	RW: GH, GM, KE, LS, MW, SD, SZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, ML, MR, NE, SN, TD, TG			

<u>AU 9885393</u>	<u>A1</u>	<u>19990104</u>	<u>AU 1998-85393</u>	<u>19980615</u>
<u>EP 996444</u>	<u>A1</u>	<u>20000503</u>	<u>EP 1998-936364</u>	<u>19980615</u>
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, FI, RO				
<u>BR 9810172</u>	A	20000808	<u>BR 1998-10172</u>	19980615
<u>JP 2002504137</u>	T2	20020205	<u>JP 1999-503755</u>	19980615
<u>ZA 9805238</u>	A	20000217	<u>ZA 1998-5238</u>	19980617
<u>NO 9906266</u>	A	19991217	<u>NO 1999-6266</u>	19991217
<u>US 2002004515</u>	A1	20020110	<u>US 2001-925394</u>	20010809
<u>US 2002137772</u>	A1	20020926	<u>US 2002-99161</u>	20020313

PRIORITY APPLN. INFO.:

<u>GB 1997-12857</u>	A	19970618
<u>GB 1998-6706</u>	A	19980327
<u>WO 1998-EP3690</u>	W	19980615
<u>US 1999-446030</u>	B1	19991215
<u>US 2001-925394</u>	B1	20010809

AB A **method** for the treatment and/or prophylaxis of **diabetes mellitus**, conditions assocd. with **diabetes mellitus**, and certain complications thereof, in a mammal which **method** comprises administering an effective nontoxic and pharmaceutically acceptable amt. of an insulin sensitizer rosiglitazone (I) and a biguanide antihyperglycemic agent such as metformin. Pharmacokinetics of I and metformin administered alone or in combination are described. Formulations for prepg. tablets contg. I is presented.

IT 155141-29-0, Rosiglitazone maleate

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(treatment of **diabetes** with thiazolidinedione insulin sensitizer and metformin)

RN 155141-29-0 HCPLUS

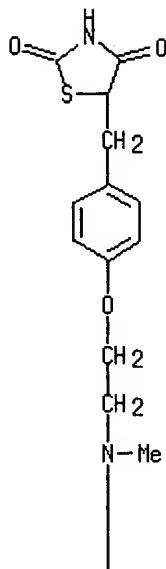
CN 2,4-Thiazolidinedione, 5-[[4-[2-(methyl-2-pyridinylamino)ethoxy]phenyl]methyl]-, (2Z)-2-butenedioate (1:1) (9CI) (CA INDEX NAME)

CM 1

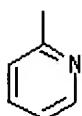
CRN 122320-73-4

CMF C18 H19 N3 O3 S

PAGE 1-A



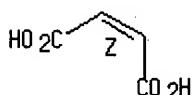
PAGE 2-A



CM 2

CRN 110-16-7
CMF C4 H4 O4

Double bond geometry as shown.



REFERENCE COUNT: 5 THERE ARE 5 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L9 ANSWER 11 OF 17 HCAPLUS COPYRIGHT 2002 ACS

Full	Citing
Text	References

ACCESSION NUMBER: 1998:764284 HCAPLUS
 DOCUMENT NUMBER: 130:10664
 TITLE: Use of 5-(4-(2-(N-methyl-N-(2-pyridyl)amino)ethoxy)benzyl)-2,4-thiazolidinedione in the treatment of polycystic ovary syndrome and gestational **diabetes**
 INVENTOR(S): Antonucci, Tammy; Lockwood, Dean; Norris, Rebecca
 PATENT ASSIGNEE(S): Warner-Lambert Company, USA
 SOURCE: PCT Int. Appl., 55 pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 7
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
<u>WO 9851305</u>	<u>A1</u>	<u>19981119</u>	<u>WO 1998-US10113</u>	<u>19980514</u>
W: AL, AU, BA, BB, BG, BR, CA, CN, CZ, EE, GE, GW, HU, ID, IL, IS, JP, KR, LC, LK, LR, LT, LV, MG, MK, MN, MX, NO, NZ, PL, RO, SG, SI, SK, SL, TR, TT, UA, US, UZ, VN, YU, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM				
RW: GH, GM, KE, LS, MW, SD, SZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, ML, MR, NE, SN, TD, TG				
<u>ZA 9804084</u>	<u>A</u>	<u>19981120</u>	<u>ZA 1998-4084</u>	<u>19980514</u>
<u>AU 9874949</u>	<u>A1</u>	<u>19981208</u>	<u>AU 1998-74949</u>	<u>19980514</u>
<u>AU 731690</u>	<u>B2</u>	<u>20010405</u>		
<u>EP 981346</u>	<u>A1</u>	<u>20000301</u>	<u>EP 1998-922391</u>	<u>19980514</u>
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO				
<u>BR 9809120</u>	<u>A</u>	<u>20000801</u>	<u>BR 1998-9120</u>	<u>19980514</u>
<u>JP 2001525827</u>	<u>T2</u>	<u>20011211</u>	<u>JP 1998-549654</u>	<u>19980514</u>
<u>AU 9952576</u>	<u>A1</u>	<u>19991202</u>	<u>AU 1999-52576</u>	<u>19991001</u>
<u>AU 749416</u>	<u>B2</u>	<u>20020627</u>		
<u>NO 9905549</u>	<u>A</u>	<u>19991112</u>	<u>NO 1999-5549</u>	<u>19991112</u>
			<u>US 1997-856987</u>	<u>A 19970515</u>
			<u>AU 1997-17709</u>	<u>A3 19970403</u>
			<u>WO 1998-US10113</u>	<u>W 19980514</u>

PRIORITY APPLN. INFO.:

AB Novel **methods** of using thiazolidinone derivs. and related antihyperglycemic agents to treat populations at risk for developing noninsulin-dependent **diabetes mellitus** (NIDDM) and complications arising therefrom are disclosed. In one embodiment, the compds. of the invention are used to treat polycystic ovary syndrome to prevent or delay the onset of noninsulin-dependent **diabetes mellitus**. In another embodiment, the compds. of the invention are used to treat gestational **diabetes** to prevent or delay the onset of noninsulin-dependent **diabetes mellitus**.

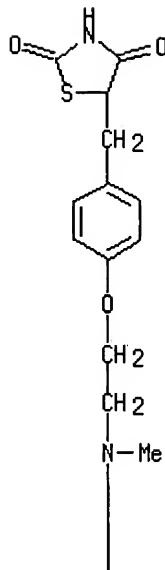
IT 122320-73-4

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
(ste ns treatment of polycystic ovary syndrome and gestational **diabetes** and prevention of NIDDM development by (methyl)pyridyl)

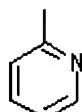
RN 122320-73-4 HCPLUS

CN 2,4-Thiazolidinedione, 5-[[4-[2-(methyl-2-pyridinylamino)ethoxy]phenyl]methyl]- (9CI) (CA INDEX NAME)

PAGE 1-A



PAGE 2-A



REFERENCE COUNT: 8 THERE ARE 8 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L9 ANSWER 12 OF 17 HCAPLUS COPYRIGHT 2002 ACS

Full Citing
 Text References

ACCESSION NUMBER: 1998:672463 HCAPLUS
 DOCUMENT NUMBER: 129:270626
 TITLE: **Methods** and compositions for treating and/or preventing non-insulin dependent **diabetes mellitus** (NIDDM) using specific retinoid compounds
 Pfahl, Magnus; Lernhardt, Waldemar; Fanjol, Andrea
 Centre International de Recherches Dermatologiques
 Galderma, Fr.
 SOURCE: PCT Int. Appl., 21 pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 1
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
<u>WO 9842340</u>	A1	19981001	<u>WO 1998-US5591</u>	19980324
W: AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE, DK, EE, ES, FI, GB, GE, GH, GM, GW, HU, ID, IL, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM				
RW: GH, GM, KE, LS, MW, SD, SZ, UG, ZW, AT, BE, CH, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, ML, MR, NE, SN, TD, TG				
<u>AU 9865763</u>	A1	19981020	<u>AU 1998-65763</u>	19980324

EP 1019049	A1	20000719	EP 1998-911919	19980324
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, FI				
BR 9808054	A	20001107	BR 1998-8054	19980324
JP 2001521551	T2	20011106	JP 1998-545851	19980324
NO 9904612	A	19991124	NO 1999-4612	19990902
<u>PRIORITY APPLN. INFO.:</u>				
US 1997-35604P P 19970324				
WO 1998-US5591 W 19980324				

AB **Methods** are provided for treating and/or preventing non-insulin dependent **diabetes mellitus** (NIDDM) in subjects having or at substantial risk of developing NIDDM, using specific retinoid compds. that are structurally related to 9-cis retinoid acid which induce the differentiation of preadipocytes into adipocytes. These compds. may be administered alone or in combination with other anti-**diabetogenic** agents such as thiazolidinediones.

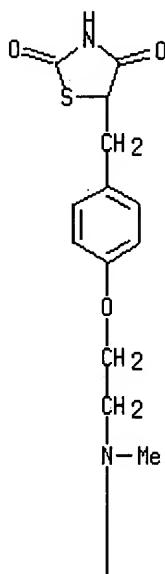
IT 122320-73-4

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
(retinoid compds. with other agents for treating and/or preventing non-insulin dependent **diabetes mellitus**)

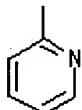
RN 122320-73-4 HCAPLUS

CN 2,4-Thiazolidinedione, 5-[[4-[2-(methyl-2-pyridinylamino)ethoxy]phenyl]methyl]-(9CI) (CA INDEX NAME)

PAGE 1-A



PAGE 2-A



L9 ANSWER 13 OF 17 HCAPLUS COPYRIGHT 2002 ACS

Full Text	Citing References
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ACCESSION NUMBER:

1998:41808 HCAPLUS

DOCUMENT NUMBER:

128:123811

TITLE:

Use of thiazolidinedione derivatives and related

antihyperglycemic agents in the treatment of insulin-resistant subjects with normal glucose tolerance in order to prevent or delay the onset of noninsulin-dependent **diabetes mellitus**

INVENTOR(S): Olefsky, Jerrold M.
 PATENT ASSIGNEE(S): Sankyo Company, Limited, Japan
 SOURCE: U.S., 16 pp.
 CODEN: USXXAM
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 1
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
<u>US 5708012</u>	<u>A</u>	<u>19980113</u>	<u>US 1995-431266</u>	<u>19950428</u>

OTHER SOURCE(S): MARPAT 128:123811

AB **Methods** are disclosed for using thiazolidinone derivs. and related antihyperglycemic agents to treat populations exhibiting insulin-resistant non-impaired glucose tolerance in order to prevent or delay the onset of noninsulin-dependent **diabetes mellitus** and complications arising therefrom. In an outpatient trial with nondiabetic, obese patients, some of whom had impaired glucose tolerance, (+)-5-[(4-[(3,4-dihydro-6-hydroxy-2,5,7,8-tetramethyl-2H-1-benzopyran-2-yl)methoxy]phenyl)methyl]-2,4-thiazolidinedione (troglitazone) normalized glucose tolerance and markedly improved insulin resistance and hyperinsulinemia.

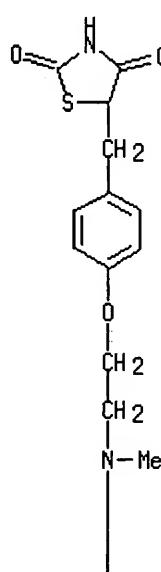
IT 122320-73-4

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
 (thiazolidinedione derivs. and related antihyperglycemic agents in treatment of insulin-resistant subjects with normal glucose tolerance to prevent or delay onset of noninsulin-dependent **diabetes mellitus**)

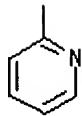
RN 122320-73-4 HCPLUS

CN 2,4-Thiazolidinedione, 5-[[4-[2-(methyl-2-pyridinylamino)ethoxy]phenyl]methyl]-(9CI) (CA INDEX NAME)

PAGE 1-A



PAGE 2-A



L9 ANSWER 14 OF 17 HCPLUS COPYRIGHT 2002 ACS

Full Text	Citing References
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ACCESSION NUMBER: 1997:329275 HCPLUS
 DOCUMENT NUMBER: 126:308792
 TITLE: Treating NIDDM with RXR agonists
 INVENTOR(S): Heyman, Richard A.; Cesario, Rosemary; Mukherjee, Ranjan
 PATENT ASSIGNEE(S): Ligand Pharmaceuticals Incorporated, USA
 SOURCE: PCT Int. Appl., 56 pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 3
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
<u>WO 9710819</u>	<u>A1</u>	<u>19970327</u>	<u>WO 1996-US14904</u>	<u>19960917</u>
W: AL, AM, AT, AU, AZ, BB, BG, BR, BY, CA, CH, CN, CZ, DE, DK, EE, ES, FI, GB, GE, HU, IS, JP, KE, KG, KP, KR, KZ, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, TJ, TM, TR, TT, UA, UG, UZ, VN, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM				
RW: KE, LS, MW, SD, SZ, UG, AT, BE, CH, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, ML				
<u>CA 2232288</u>	<u>AA</u>	<u>19970327</u>	<u>CA 1996-2232288</u>	<u>19960917</u>
<u>AU 9670742</u>	<u>A1</u>	<u>19970409</u>	<u>AU 1996-70742</u>	<u>19960917</u>
<u>AU 725998</u>	<u>B2</u>	<u>20001026</u>		
<u>EP 859608</u>	<u>A1</u>	<u>19980826</u>	<u>EP 1996-931613</u>	<u>19960917</u>
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, FI				
<u>BR 9610624</u>	<u>A</u>	<u>19990316</u>	<u>BR 1996-10624</u>	<u>19960917</u>
<u>JP 11511472</u>	<u>T2</u>	<u>19991005</u>	<u>JP 1996-512842</u>	<u>19960917</u>
<u>US 6028052</u>	<u>A</u>	<u>20000222</u>	<u>US 1996-710309</u>	<u>19960917</u>
<u>US 5972881</u>	<u>A</u>	<u>19991026</u>	<u>US 1997-979725</u>	<u>19971126</u>
<u>NO 9801192</u>	<u>A</u>	<u>19980518</u>	<u>NO 1998-1192</u>	<u>19980317</u>
<u>US 6228862</u>	<u>B1</u>	<u>20010508</u>	<u>US 1999-309370</u>	<u>19990511</u>
<u>US 6316404</u>	<u>B1</u>	<u>20011113</u>	<u>US 2000-745681</u>	<u>20001222</u>
<u>PRIORITY APPLN. INFO.:</u>			<u>US 1995-3869P</u>	<u>P</u> <u>19950918</u>
			<u>US 1995-4897P</u>	<u>P</u> <u>19951006</u>
			<u>US 1996-9884P</u>	<u>P</u> <u>19960110</u>
			<u>US 1996-18318P</u>	<u>P</u> <u>19960524</u>
			<u>US 1996-21839P</u>	<u>P</u> <u>19960710</u>
			<u>US 1996-710309</u>	<u>B3</u> <u>19960917</u>
			<u>WO 1996-US14904</u>	<u>W</u> <u>19960917</u>
			<u>US 1997-979725</u>	<u>A1</u> <u>19971126</u>
			<u>US 1999-309370</u>	<u>A3</u> <u>19990511</u>

AB This invention relates to methods and compns. for the treatment of non-insulin-dependent diabetes mellitus using an RXR agonist alone or in combination with a PPAR γ agonist such as thiazolidinedione compd. Example RXR agonists are LGD 1069, ALRT 1957 and LG 100268.

IT 122320-73-4, BRL 49653

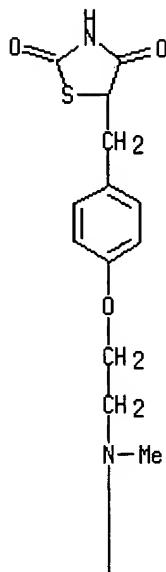
RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
 (noninsulin dependent diabetes treatment with RXR agonists)

RN 122320-73-4 HCPLUS

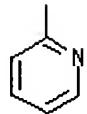
CN 2,4-Thiazolidinedione, 5-[[4-[2-(methyl-2-pyridinylamino)ethoxy]phenyl]met

hyl]- (9CI) (CA INDEX NAME)

PAGE 1-A



PAGE 2-A



L9 ANSWER 15 OF 17 HCAPLUS COPYRIGHT 2002 ACS

 Full Citing
 Text References

ACCESSION NUMBER: 1997:231131 HCAPLUS
 DOCUMENT NUMBER: 126:207528
 TITLE: A thiazolidione derivative for reducing the amount of exogenous insulin administered to a patient having noninsulin-dependent **diabetes mellitus**
 INVENTOR(S): Whitcomb, Randall W.
 PATENT ASSIGNEE(S): Warner-Lambert Company, USA; Whitcomb, Randall W.
 SOURCE: PCT Int. Appl., 43 pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 1
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
<u>WO 9705875</u>	<u>A2</u>	<u>19970220</u>	<u>WO 1996-US12430</u>	<u>19960729</u>
<u>WO 9705875</u>	<u>A3</u>	<u>19970327</u>		
W: AU, BG, CA, CN, CZ, EE, GE, HU, IL, JP, KR, LT, LV, MX, NO, NZ, PL, RO, SG, SI, SK, UA, US, UZ, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM RW: AT, BE, CH, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE				
<u>CA 2221241</u>	<u>AA</u>	<u>19970220</u>	<u>CA 1996-2221241</u>	<u>19960729</u>
<u>AU 9666411</u>	<u>A1</u>	<u>19970305</u>	<u>AU 1996-66411</u>	<u>19960729</u>
<u>AU 724989</u>	<u>B2</u>	<u>20001005</u>		
<u>EP 851757</u>	<u>A2</u>	<u>19980708</u>	<u>EP 1996-926171</u>	<u>19960729</u>
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,				

IE, SI, LT, LV, FI

<u>CN 1192683</u>	<u>A</u>	<u>19980909</u>	<u>CN 1996-196191</u>	<u>19960729</u>
<u>JP 11510508</u>	<u>T2</u>	<u>19990914</u>	<u>JP 1996-508479</u>	<u>19960729</u>
<u>NO 9800556</u>	<u>A</u>	<u>19980209</u>	<u>NO 1998-556</u>	<u>19980209</u>
<u>PRIORITY APPLN. INFO.:</u>			<u>US 1995-2098P</u>	<u>P 19950810</u>
			<u>WO 1996-US12430</u>	<u>W 19960729</u>

OTHER SOURCE(S): MARPAT 126:207528

AB This invention provides a **method** of reducing the amt. of exogenous insulin administered to a patient having noninsulin-dependent **diabetes mellitus** by administering to a patient a therapeutically effective amt. of a thiazolidione deriv. and/or a related compd. Seventeen patients with noninsulin-dependent **diabetes mellitus** that were still on insulin were treated with thiazolidinedione deriv. (400 mg/day) for 8 wk. Ten patients have had a mean decrease of 45% (39 units) in their daily dose of insulin and appear to be continuing to reduce their insulin requirements. At the same time, their glycemic control was improving with a mean decrease of 15% (36 mg/dL) in blood glucose. A total of 7 patients have had their insulin discontinued after 8 wk.

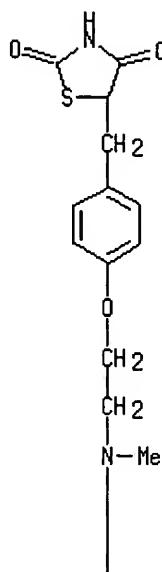
IT 122320-73-4, BRL 49653

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
(thiazolidione deriv. and/or related compds. for reducing amt. of exogenous insulin in humans with noninsulin-dependent **diabetes mellitus**)

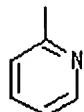
RN 122320-73-4 HCPLUS

CN 2,4-Thiazolidinedione, 5-[[4-[2-(methyl-2-pyridinylamino)ethoxy]phenyl]methyl]-(9CI) (CA INDEX NAME)

PAGE 1-A



PAGE 2-A



Full Text	Citing References
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ACCESSION NUMBER: 1996:713048 HCAPLUS
 DOCUMENT NUMBER: 125:319877
 TITLE: Adipocyte containing ob gene promoter for screening modulators useful in treatment of anorexia, obesity, and other diseases
 INVENTOR(S): Briggs, Michael R.; Auwerx, Johan; De Vos, Piet; Staels, Bart; Croston, Glenn E.; Miller, Stephen G.
 PATENT ASSIGNEE(S): Ligand Pharmaceuticals Incorporated, USA; Institut Pasteur De Lille
 SOURCE: PCT Int. Appl., 166 pp.
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 2
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
<u>WO 9629405</u>	<u>A2</u>	<u>19960926</u>	<u>WO 1996-US3808</u>	<u>19960319</u>
<u>WO 9629405</u>	<u>A3</u>	<u>19961128</u>		
W: AL, AM, AT, AU, AZ, BB, BG, BR, BY, CA, CH, CN, CZ, DE, DK, EE, ES, FI, GB, GE, HU, IS, JP, KE, KG, KP, KR, KZ, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI				
RW: KE, LS, MW, SD, SZ, UG, AT, BE, CH, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, ML				
<u>CA 2215387</u>	<u>AA</u>	<u>19960926</u>	<u>CA 1996-2215387</u>	<u>19960319</u>
<u>AU 9655248</u>	<u>A1</u>	<u>19961008</u>	<u>AU 1996-55248</u>	<u>19960319</u>
<u>EP 815230</u>	<u>A2</u>	<u>19980107</u>	<u>EP 1996-912428</u>	<u>19960319</u>
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, FI				

PRIORITY APPLN. INFO.:

<u>US 1995-408584</u>	<u>A</u>	<u>19950320</u>
<u>US 1995-418096</u>	<u>A</u>	<u>19950405</u>
<u>US 1995-510584</u>	<u>A</u>	<u>19950802</u>
<u>US 1995-558588</u>	<u>A</u>	<u>19951030</u>
<u>US 1995-7390P</u>	<u>P</u>	<u>19951121</u>
<u>US 1995-7721P</u>	<u>P</u>	<u>19951130</u>
<u>US 1995-8601P</u>	<u>P</u>	<u>19951214</u>
<u>WO 1996-US3808</u>	<u>W</u>	<u>19960319</u>

AB This invention relates to the isolation and cloning of the promoter and other control regions of a human ob gene. It provides a **method** for identifying and screening for agents useful for the treatment of diseases and pathol. conditions affected by the level of expression of an ob gene. These agents interact directly or indirectly with the promoter or other control regions of the ob gene. A PPAR γ agonist, BRL49653, has been identified to be useful in treating anorexia, cachexia, and other diseases characterized by insufficient food intake or body wt. loss. Modulators of ob gene expression may be used to treat other diseases such as obesity, diabetes, hypertension, cardiovascular diseases and infertility.

IT 122320-73-4, BRL49653

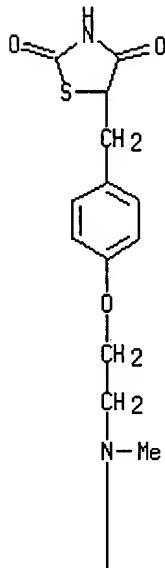
RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(PPAR γ agonist; adipocyte contg. ob gene promoter for screening modulators useful in treatment of anorexia, obesity, and other diseases)

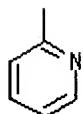
RN 122320-73-4 HCAPLUS

CN 2,4-Thiazolidinedione, 5-[[4-[2-(methyl-2-pyridinylamino)ethoxy]phenyl]methyl]- (9CI) (CA INDEX NAME)

PAGE 1-A



PAGE 2-A



L9 ANSWER 17 OF 17 HCPLUS COPYRIGHT 2002 ACS

Full Text	Citing References
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ACCESSION NUMBER: 1996:71475 HCPLUS
 DOCUMENT NUMBER: 124:106679
 TITLE: Thiazolidinedione derivatives and related antihyperglycemic agents in the treatment of impaired glucose tolerance to prevent or delay the onset of noninsulin-dependent **diabetes mellitus**
 INVENTOR(S): Olefsky, Jerrold; Antonucci, Tammy; Lockwood, Dean; Norris, Rebecca
 PATENT ASSIGNEE(S): Sankyo Co., Ltd., Japan
 SOURCE: U.S., 15 pp. Cont.-in-part of U.S. Ser. No. 122,251, abandoned.
 CODEN: USXXAM
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 7
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
<u>US 5478852</u>	A	19951226	<u>US 1994-293899</u>	19940823
<u>US 5478852</u>	C1	20010313		
<u>WO 9507697</u>	A2	19950323	<u>WO 1994-US10187</u>	19940909
<u>WO 9507697</u>	A3	19950511		
W: AU, CA, CN, CZ, FI, HU, JP, KR, NO, NZ, RU, SK RW: AT, BE, CH, DE, DK, ES, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE				
<u>WO 9507694</u>	A1	19950323	<u>WO 1994-US10389</u>	19940914
W: AU, CA, CN, CZ, FI, HU, JP, KR, MW, NO, NZ, RU RW: AT, BE, CH, DE, DK, ES, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE				
<u>AU 9478351</u>	A1	19950403	<u>AU 1994-78351</u>	19940914

AU 679572	B2	19970703		
<u>EP 719140</u>	<u>A1</u>	<u>19960703</u>	<u>EP 1994-929204</u>	<u>19940914</u>
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CN 1134669	A	19961030	CN 1994-194058	19940914
JP 09502727	T2	19970318	JP 1995-509333	19940914
JP 3081245	B2	20000828		
HU 75874	A2	19970528	HU 1996-653	19940914
CZ 283207	B6	19980114	CZ 1996-2822	19940914
CZ 283208	B6	19980114	CZ 1996-2823	19940914
CZ 283339	B6	19980318	CZ 1996-793	19940914
JP 2000239167	A2	20000905	JP 2000-71978	19940914
JP 2000273043	A2	20001003	JP 2000-71977	19940914
NO 9601041	A	19960514	NO 1996-1041	19960314
FI 9601213	A	19960514	FI 1996-1213	19960315
AU 9717709	A1	19970529	AU 1997-17709	19970403
AU 706947	B2	19990701		
AU 9717710	A1	19970529	AU 1997-17710	19970403
AU 9952576	A1	19991202	AU 1999-52576	19991001
AU 749416	B2	20020627		
NO 2000002963	A	20000609	NO 2000-2963	20000609
NO 2000002964	A	20000609	NO 2000-2964	20000609
<u>PRIORITY APPLN. INFO.:</u>				
		US 1993-122251	B2	19930915
		US 1994-292585	A	19940823
		US 1994-293899		19940823
		JP 1994-509333	A3	19940914
		JP 1995-509333	A3	19940914
		WO 1994-US10389	W	19940914
		AU 1997-17709	A3	19970403

OTHER SOURCE(S): MARPAT 124:106679

AB Novel methods of using thiazolidinone derivs. and related antihyperglycemic agents to treat populations experiencing impaired glucose tolerance in order to prevent or delay the onset of noninsulin-dependent **diabetes mellitus** and complications arising therefrom, are disclosed. Effects of (+)-5-[[4-[(3,4-dihydro-6-hydroxy-2,5,7,8-tetramethyl-2H-1-benzopyran-2-yl)methoxy]phenyl]methyl]-2,4-thiazolidinedione (troglitazone) was clin. tested with patients with impaired glucose tolerance by the WHO criteria; the results showed that treatment with troglitazone correlated to redn. of fasting insulin levels and return of glucose tolerance to the normal range for ~70% of the subjects.

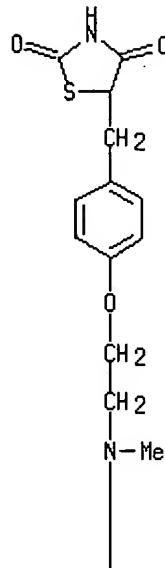
IT 122320-73-4

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
(thiazolidinedione derivs. in prevention of onset of noninsulin-dependent **diabetes**)

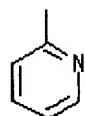
RN 122320-73-4 HCAPLUS

CN 2,4-Thiazolidinedione, 5-[[4-[2-(methyl-2-pyridinylamino)ethoxy]phenyl]methyl]- (9CI) (CA INDEX NAME)

PAGE 1-A



PAGE 2-A



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 L6 92 S L5 AND METHOD
 L7 56 S L6 AND DIAB?
 L8 44 S L7 AND MELLIT?
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11 BLACKLER, P?/AU
 L11 3 L10 AND BLACKLER, P?/AU

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L11 ANSWER 1 OF 3 HCAPLUS COPYRIGHT 2002 ACS

Full Text	Citing References
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ACCESSION NUMBER: 2000:772629 HCPLUS
 DOCUMENT NUMBER: 133:340315
 TITLE: Therapeutic action and properties of a **polymorphic** form of 5-[4-[2-(N-methyl-N-(2-pyridyl)amino)ethoxy]benzyl]thiazolidine-2,4-dione, maleic acid salt
 INVENTOR(S): **Blackler, Paul David James**; Browne, Christine Marie; Coakley, Timothy G.; Giles, Robert Gordon; Morrissey, Gillian
 PATENT ASSIGNEE(S): SmithKline Beecham PLC, UK; SmithKline Beecham (Cork) Limited
 SOURCE: PCT Int. Appl., 21 pp.
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 1
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
<u>WO 2000064896</u>	A1	20001102	<u>WO 2000-GB1520</u>	20000419
W: AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, LZ, LC, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM				
RW: GH, GM, KE, LS, MW, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG				
<u>EP 1173435</u>	A1	20020123	<u>EP 2000-920892</u>	20000419
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO				
<u>BR 2000009932</u>	A	20020409	<u>BR 2000-9932</u>	20000419
<u>NO 2001005147</u>	A	20011217	<u>NO 2001-5147</u>	20011022
<u>PRIORITY APPLN. INFO.:</u>				
		GB 1999-9473	A	19990423
		GB 1999-12196	A	19990525
		<u>WO 2000-GB1520</u>	W	20000419

AB A **polymorphic** form of 5-[4-[2-(N-methyl-N-(2-pyridyl)amino)ethoxy]benzyl]thiazolidine-2, 4-dione, maleic acid salt (the "Polymorph") characterized in that it provides: (i) an IR spectrum contg. peaks at 1763, 912, 856 and 709 cm⁻¹; and/or (ii) a Raman spectrum contg. peaks at 1762, 1284, 912 and 888 cm⁻¹; and/or (iii) a solid-state ¹³C NMR spectrum contg. peaks at 111.0, 113.6, 119.8, 129.1, 130.9, 131.8, 134.7, 138.7, 146.5, 152.7, 157.5, 169.5, 171.0, 178.7 ppm; and/or (iv) an x-ray powder diffraction (XRPD) pattern which gives calcd. lattice spacings at 5.87, 5.30, 4.69, 4.09, 3.88, 3.61, 3.53 and 3.46 Angstroms; a process for prepg. such a compd., a pharmaceutical compn. contg. such a compd. and the use of such a compd. in medicine.

IT 155141-29-0

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); PRP (Properties); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
 (antidiabetic action and properties of **polymorphic** form of [(N-methyl-N-(pyridyl)amino)ethoxy]benzyl]thiazolidinedione maleate)

RN 155141-29-0 HCPLUS

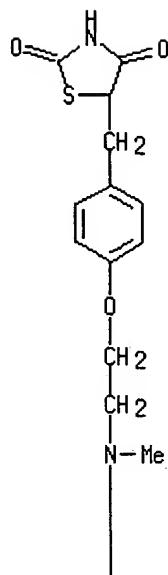
CN 2,4-Thiazolidinedione, 5-[(4-[2-(methyl-2-pyridinylamino)ethoxy]phenyl)methyl]-, (2Z)-2-butenedioate (1:1) (9CI) (CA INDEX NAME)

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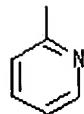
CRN 122320-73-4

CMF C18 H19 N3 O3 S

PAGE 1-A



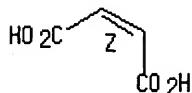
PAGE 2-A



CM 2

CRN 110-16-7
CMF C4 H4 O4

Double bond geometry as shown.



REFERENCE COUNT:

4

THERE ARE 4 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L11 ANSWER 2 OF 3

HCAPLUS COPYRIGHT 2002 ACS

Full Text	Citing References
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ACCESSION NUMBER:

2000:772627 HCAPLUS

DOCUMENT NUMBER:

133:340314

TITLE:

Therapeutic action and properties of a **polymorphic** form of 5-[4-[2-(N-methyl-N-(2-pyridyl)amino)ethoxy]benzyl]thiazolidine-2,4-dione, maleic acid salt

INVENTOR(S):

Blackler, Paul David James; Giles, Robert Gordon; Moore, Stephen; Sasse, Michael John

PATENT ASSIGNEE(S):

SmithKline Beecham PLC, UK

SOURCE:

PCT Int. Appl., 19 pp.

DOCUMENT TYPE:

CODEN: PIXXD2

LANGUAGE:

Patent

English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2000064893	A2	20001102	WO 2000-GB1522	20000419
WO 2000064893	A3	20010125		
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM				
RW: GH, GM, KE, LS, MW, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG				
EP 1175418	A2	20020130	EP 2000-922793	20000419
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO				
BR 2000009935	A	20020416	BR 2000-9935	20000419
NO 2001005148	A	20011217	NO 2001-5148	20011022
<u>PRIORITY APPLN. INFO.:</u>				
		GB 1999-9471	A	19990423
		GB 1999-12195	A	19990525
		WO 2000-GB1522	W	20000419

AB A **polymorphic** form of 5-[4-[2-(N-methyl-N-(2-pyridyl)amino)ethoxy]benzyl]thiazolidine-2,4-dione, maleic acid salt (the "Polymorph") characterized in that it provides: (i) an infra red spectrum contg. peaks at 1752, 1546, 1154, 621, and 602 cm^{-1} ; and/or (ii) a Raman spectrum contg. peaks at 1751, 1243 and 602 cm^{-1} ; and/or (iii) a solid-state NMR spectrum contg. peaks at 111.9, 114.8, 119.6, 129.2, 134.0, 138.0, 144.7, 153.2, 157.1, 170.7, 172.0 and 175.0 ppm; and/or (iv) an x-ray powder diffraction (XRPD) pattern which gives calcd. lattice spacings of 6.46, 5.39, 4.83, 4.68, 3.71, 3.63, 3.58, and 3.48 Angstroms; a process for prep. such a compd., a pharmaceutical compn. contg. such a compd. and the use of such a compd. in medicine.

IT 168553-12-6

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); PRP (Properties); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(antidiabetic action of **polymorphic** form of
[(N-methyl-N-(pyridyl)amino)ethoxy]benzyl]thiazolidinedione maleate)

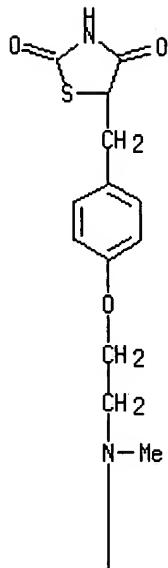
RN 168553-12-6 HCPLUS

CN 2,4-Thiazolidinedione, 5-[[4-[2-(methyl-2-pyridinylamino)ethoxy]phenyl]methyl]-, (2Z)-2-butenedioate (9CI) (CA INDEX NAME)

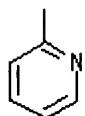
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PAGE 1-A



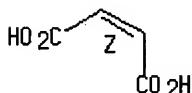
PAGE 2-A



CM 2

CRN 110-16-7
CMF C4 H4 O4

Double bond geometry as shown.



L11 ANSWER 3 OF 3 HCAPLUS COPYRIGHT 2002 ACS

[Full Text](#) [Citing References](#)

ACCESSION NUMBER: 2000:772626 HCAPLUS
 DOCUMENT NUMBER: 133:340313
 TITLE: Therapeutic action and properties of a **polymorphic** form of 5-[4-[2-(N-methyl-N-(2-pyridyl)amino)ethoxy]benzyl]thiazolidine-2,4-dione, maleic acid salt
 INVENTOR(S): Blackler, Paul David James; Giles, Robert Gordon; Sasse, Michael John
 PATENT ASSIGNEE(S): SmithKline Beecham P.L.C., UK
 SOURCE: PCT Int. Appl., 18 pp.
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 1
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2000064892	A2	20001102	WO 2000-GB1514	20000419
WO 2000064892	A3	20010125		
W: AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM				
RW: GH, GM, KE, LS, MW, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG				
EP 1173434	A2	20020123	EP 2000-920889	20000419
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO				
BR 2000009934	A	20020604	BR 2000-9934	20000419
NO 2001005149	A	20011217	NO 2001-5149	20011022
PRIORITY APPLN. INFO.:				
GB 1999-9472 A 19990423				
GB 1999-12197 A 19990525				
WO 2000-GB1514 W 20000419				

AB A **polymorphic** form of 5-[4-[2-(N-methyl-N-(2-pyridyl)amino)ethoxy]benzyl]thiazolidine-2,4-dione, maleic acid salt (the "Polymorph") characterized in that it: (i) provides an IR spectrum contg. peaks at 1360, 1326, 1241, 714 and 669 cm⁻¹; and/or (ii) provides a Raman spectrum contg. peaks at 1581, 768, 670, 271 and 226 cm⁻¹; and/or (iii) provides a solid-state NMR spectrum contg. peaks at chem. shifts substantially; and/or (iv) provides an x-ray powder diffraction (XRPD) pattern contg. peaks; a process for prep. such a compd., a pharmaceutical compn. contg. such a compd. and the use of such a compd. in medicine.

IT 168553-12-6

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); PRP (Properties); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(antidiabetic action of **polymorphic** form of
[(N-methyl-N-(pyridyl)amino)ethoxy]benzyl]thiazolidinedione maleate)

RN 168553-12-6 HCPLUS

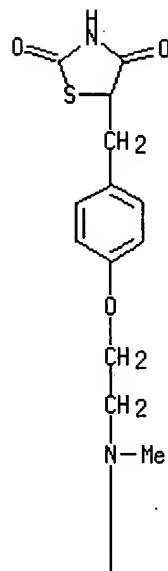
CN 2,4-Thiazolidinedione, 5-[[4-[2-(methyl-2-pyridinylamino)ethoxy]phenyl]methyl]-, (2Z)-2-butenedioate (9CI) (CA INDEX NAME)

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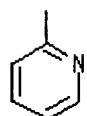
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CMF C18 H19 N3 O3 S

PAGE 1-A



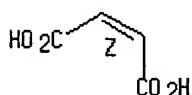
PAGE 2-A



CM 2

CRN 110-16-7
CMF C4 H4 O4

Double bond geometry as shown.



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 L3 48 S L1 FULL

FILE 'HCAPLUS' ENTERED AT 20:14:30 ON 03 OCT 2002

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 L5 324 S L3/THU
 L6 92 S L5 AND METHOD
 L7 56 S L6 AND DIAB?
 L8 44 S L7 AND MELLIT?
 L9 17 S L8 AND PD < MAY 20 1999
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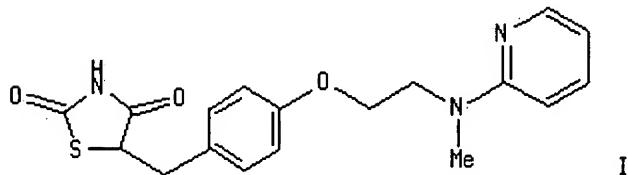
L12 ANSWER 1 OF 4 HCAPLUS COPYRIGHT 2002 ACS

Full Citing
 Text References

ACCESSION NUMBER: 2002:504785 HCAPLUS
 DOCUMENT NUMBER: 137:83621
 TITLE: Preparation and use of 5-[4-[2-(N-methyl-N-(2-pyridyl)amino)ethoxy]benzyl]thiazolidine-2,4-dione methanesulfonate
 INVENTOR(S): Craig, Andrew Simon; Ho, Tim Chien Ting; Millan, Michael; O'Keefe, Deirdre
 PATENT ASSIGNEE(S): Smithkline Beecham P.L.C., UK
 SOURCE: PCT Int. Appl., 41 pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 1
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2002051839	A1	20020704	WO 2001-GB5751	20011221
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM				
RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG				
<u>PRIORITY APPLN. INFO.:</u>			GB 2000-31521	A 20001222
			GB 2000-31524	A 20001222
			GB 2000-31526	A 20001222
			GB 2000-31528	A 20001222

GI



AB A compd. 5-[4-[2-(N-methyl-N-(2-pyridyl)amino)ethoxy]benzyl]thiazolidine-2,4-dione (I) methanesulfonate salt (II) or solvate thereof; a process for prep. I, a compn. comprising I and its therapeutic use is disclosed. Four **polymorphic** forms were prep'd. and characterized. For instance, MsOH (0.54 mL) was added to a mixt. of I (3.0 g) in EtOAc (60 mL) and was heated with agitation to reflux to give a suspension. The resulting mixt. was cooled to 21°C, the solid formed collected by filtration, washed with EtOAc and dried under vacuum for 16 h (3.73 g yield). **Polymorphic** forms I-IV were characterized by at least one of the following means: aq. solv., m.p., 1H-NMR (soln.), 13C-NMR (solid state), IR/Raman spectra, XRPD and DSC. II is a stable solid with good water solv., desirable flow properties and is amenable to large scale processing (milling). II is useful for the prevention/treatment of diabetes mellitus.

IT **439902-56-4P**

RL: IMF (Industrial manufacture); PRP (Properties); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

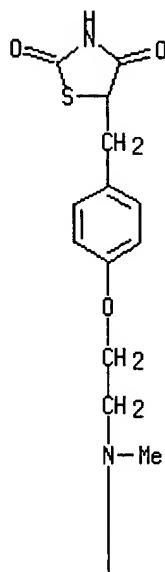
(**polymorphic** forms I-IV characterized; prepn. and characterization of 5-[4-[2-(N-Me-N-(2-pyridyl)amino)ethoxy]benzyl]thiazolidine-2,4-dione methanesulfonate)

RN **439902-56-4** HCPLUSCN **2,4-Thiazolidinedione, 5-[[4-[2-(methyl-2-pyridinylamino)ethoxy]phenyl]methyl]-, monomethanesulfonate (9CI)** (CA INDEX NAME)

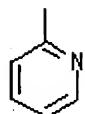
CM 1

CRN **122320-73-4**CMF **C₁₈ H₁₉ N₃ O₃ S**

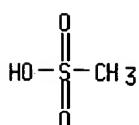
PAGE 1-A



PAGE 2-A



CM 2

CRN **75-75-2**CMF **C H₄ O₃ S**

REFERENCE COUNT:

2

THERE ARE 2 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

Full Citing
Text References

ACCESSION NUMBER: 2002:256258 HCAPLUS
 DOCUMENT NUMBER: 136:299688
 TITLE: Novel **polymorphic** forms of 5-[4-[2-[N-methyl-N-(2-pyridyl)amino]ethoxy]benzyl] thiazolidine-2,4-dione maleate and process for their preparation
 INVENTOR(S): Chebiyyam, Prabhakar; Mamillapalli, Ramabhadra Sarma; Krishnamurthi, Vyas; Seella, Vishnuvardhan Reddy; Gaddam, Om Reddy
 PATENT ASSIGNEE(S): Reddy's Research Foundation, India; Cord, Janet I.
 SOURCE: PCT Int. Appl., 40 pp.
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 1
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2002026737	A1	20020404	WO 2001-US29896	20010925
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PH, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG				
AU 2001091232	A5	20020408	AU 2001-91232	20010925
<u>PRIORITY APPLN. INFO.:</u>			IN 2000-MA805	A 20000926
			WO 2001-US29896	W 20010925

AB This invention relates to novel **polymorphic/pseudopolymorphic** forms of 5-[4-[2-[N-methyl-N-(2-pyridyl)amino]ethoxy]benzyl] thiazolidine-2,4-dione maleate (I). The invention also relates to a pharmaceutical compn. comprising the novel **polymorphic** form or their mixt. and a pharmaceutically acceptable carrier. The **polymorphic** forms of the present invention are more active, as antidiabetic agent, than the hitherto known 5-[4-[2-[N-2-methyl-N-(2-pyridyl)amino]ethoxy]benzyl] thiazolidine-2,4-dione maleate. I was dissolved in ethanol and was allowed to cool to room temp. over a period of 18 h to yield 80% of >99% pure **polymorphic** form of I.

IT 155141-29-0P

RL: PRP (Properties); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)
 (novel **polymorphic** forms of triazolidinedione maleate and process for their prepn.)

RN 155141-29-0 HCAPLUS

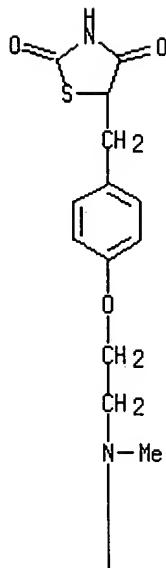
CN 2,4-Thiazolidinedione, 5-[[4-[2-(methyl-2-pyridinylamino)ethoxy]phenyl]met hyl]-, (2Z)-2-butenedioate (1:1) (9CI) (CA INDEX NAME)

CM 1

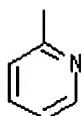
CRN 122320-73-4

CMF C18 H19 N3 O3 S

PAGE 1-A



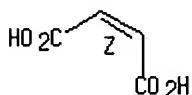
PAGE 2-A



CM 2

CRN 110-16-7
CMF C4 H4 O4

Double bond geometry as shown.



REFERENCE COUNT: 6 THERE ARE 6 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L12 ANSWER 3 OF 4 HCAPLUS COPYRIGHT 2002 ACS

Full Text	Citing References
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ACCESSION NUMBER: 2001:724803 HCAPLUS
 DOCUMENT NUMBER: 136:79548
 TITLE: Inhibition of RXR and PPAR γ ameliorates diet-induced obesity and type 2 diabetes
 AUTHOR(S): Yamauchi, Toshimasa; Waki, Hironori; Kamon, Junji; Murakami, Koji; Motojima, Kiyoto; Komeda, Kajuro; Miki, Hiroshi; Kubota, Naoto; Terauchi, Yasuo; Tsuchida, Atsuko; Tsuboyama-Kasaoka, Nobuyo; Yamauchi, Naoko; Ide, Tomohiro; Hori, Wataru; Kato, Shigeaki; Fukayama, Masashi; Akanuma, Yasuo; Ezaki, Osamu; Itai, Akiko; Nagai, Ryozo; Kimura, Satoshi; Tobe, Kazuyuki; Kagechika, Hiroyuki; Shudo, Koichi; Kadokawa, Takashi
 CORPORATE SOURCE: Department of Internal Medicine, Graduate School of Medicine, University of Tokyo, Tokyo, 113-8655, Japan

SOURCE: Journal of Clinical Investigation (2001), 108(7), 1001-1013
 CODEN: JCINAO; ISSN: 0021-9738

PUBLISHER: American Society for Clinical Investigation

DOCUMENT TYPE: Journal

LANGUAGE: English

AB PPAR γ is a ligand-activated transcription factor and functions as a heterodimer with a retinoid X receptor (RXR). Supraphysiolog. activation of PPAR γ by thiazolidinediones can reduce insulin resistance and hyperglycemia in type 2 diabetes, but these drugs can also cause wt. gain. Quite unexpectedly, a moderate redn. of PPAR γ activity obsd. in heterozygous PPAR γ -deficient mice or the Pro12Ala polymorphism in human PPAR γ , has been shown to prevent insulin resistance and obesity induced by a high-fat diet. In this study, we investigated whether functional antagonism toward PPAR γ /RXR could be used to treat obesity and type 2 diabetes. We show herein that an RXR antagonist and a PPAR γ antagonist decrease triglyceride (TG) content in white adipose tissue, skeletal muscle, and liver. These inhibitors potentiated leptin's effects and increased fatty acid combustion and energy dissipation, thereby ameliorating HF diet-induced obesity and insulin resistance. Paradoxically, treatment of heterozygous PPAR γ -deficient mice with an RXR antagonist or a PPAR γ antagonist depletes white adipose tissue and markedly decreases leptin levels and energy dissipation, which increases TG content in skeletal muscle and the liver, thereby leading to the re-emergence of insulin resistance. Our data suggested that appropriate functional antagonism of PPAR γ /RXR may be a logical approach to protection against obesity and related diseases such as type 2 diabetes.

IT 122320-73-4, Rosiglitazone

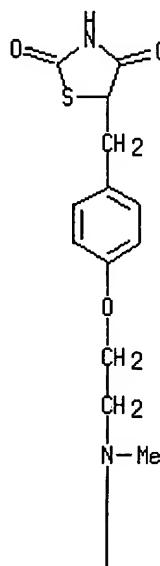
RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(inhibition of RXR and PPAR γ ameliorates diet-induced obesity and type 2 diabetes)

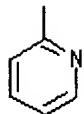
RN 122320-73-4 HCPLUS

CN 2,4-Thiazolidinedione, 5-[[4-[2-(methyl-2-pyridinylamino)ethoxy]phenyl]methyl]- (9CI) (CA INDEX NAME)

PAGE 1-A



PAGE 2-A



REFERENCE COUNT: 55 THERE ARE 55 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L12 ANSWER 4 OF 4 HCAPLUS COPYRIGHT 2002 ACS

Full Citing References
 Text

ACCESSION NUMBER: 1999:437778 HCAPLUS
 DOCUMENT NUMBER: 131:197757
 TITLE: Loss-of-function mutations in PPAR γ associated with human colon cancer
 AUTHOR(S): Sarraf, Pasha; Mueller, Elisabetta; Smith, Wendy M.; Wright, Harold M.; Kum, Jennifer B.; Aaltonen, Lauri A.; De la Chapelle, Albert; Spiegelman, Bruce M.; Eng, Charis
 CORPORATE SOURCE: Department of Cancer Biology Dana-Farber Cancer Institute Department of Cell Biology, Harvard Medical School, Boston, MA, 02115, USA
 SOURCE: Molecular Cell (1999), 3(6), 799-804
 CODEN: MOCEFL; ISSN: 1097-2765

PUBLISHER: Cell Press
 DOCUMENT TYPE: Journal
 LANGUAGE: English

AB The gamma isoform of the peroxisome proliferator-activated receptor, PPAR γ , regulates adipocyte differentiation and has recently been shown to be expressed in neoplasia of the colon and other tissues. The authors have found four somatic PPAR γ mutations among 55 sporadic colon cancers: one nonsense, one frameshift, and two missense mutations. Each greatly impaired the function of the protein. C.472delA results in deletion of the entire ligand binding domain. Q286P and K319X retain a total or partial ligand binding domain but lose the ability to activate transcription through a failure to bind to ligands. R288H showed a normal response to synthetic ligands but greatly decreased transcription and binding when exposed to natural ligands. These data indicate that colon cancer in humans is assocd. with loss-of-function mutations in PPAR γ .

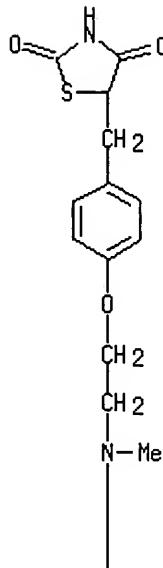
IT 122320-73-4, BRL 49653

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); BUU (Biological use, unclassified); BIOL (Biological study); USES (Uses)
 (loss-of-function mutated PPAR γ assocd. with human colon cancer binding of and transactivation response to)

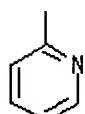
RN 122320-73-4 HCAPLUS

CN 2,4-Thiazolidinedione, 5-[[4-[2-(methyl-2-pyridinylamino)ethoxy]phenyl]methyl]- (9CI) (CA INDEX NAME)

PAGE 1-A



PAGE 2-A



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28

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 FILE LAST UPDATED: 01 May 1997 (19970501/UP)

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L3 48 S L1 FULL

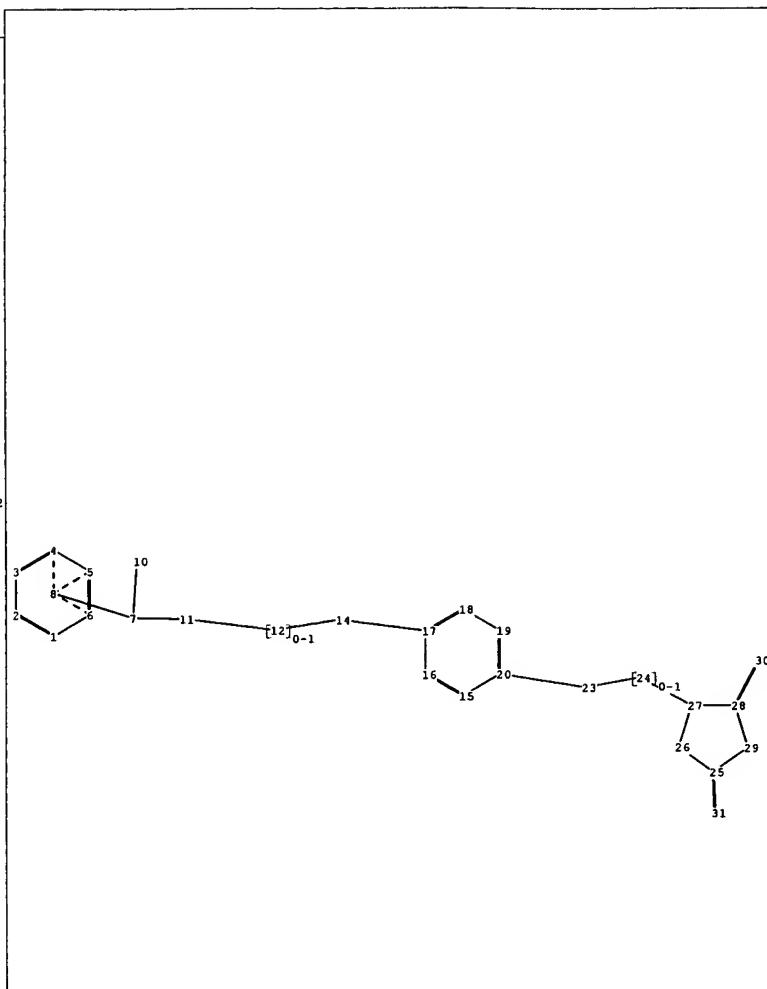
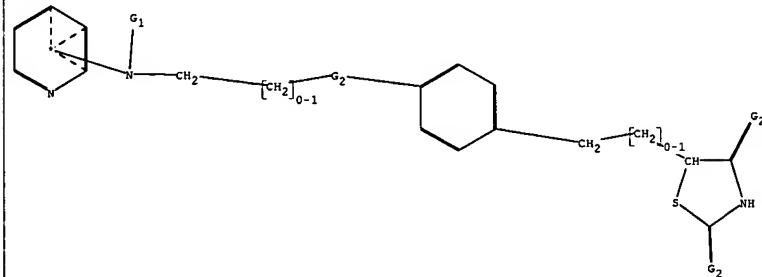
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L5 324 S L3/THU
L6 92 S L5 AND METHOD
L7 56 S L6 AND DIAB? ;
L8 44 S L7 AND MELLIT?
L9 17 S L8 AND PD < MAY 20 1999
L10 7 S L3 AND POLYMO?
L11 3 S L10 AND BLACKLER, P?/AU
L12 4 S L10 NOT L11

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L13 0 L3

=>



chain nodes :
 7 10 11 12 14 23 24 30 31

ring nodes :
 1 2 3 4 5 6 15 16 17 18 19 20 25 26 27 28 29

chain bonds :
 7-10 7-11 11-12 12-14 14-17 20-23 23-24 24-27 25-31 28-30

ring bonds :
 1-2 1-6 2-3 3-4 4-5 5-6 15-16 15-20 16-17 17-18 18-19 19-20 25-26 25-29
 26-27 27-28 28-29

exact/norm bonds :
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exact bonds :
 7-11 11-12 20-23 23-24 24-27 25-26 26-27 27-28

normalized bonds :
 1-2 1-6 2-3 3-4 4-5 5-6 15-16 15-20 16-17 17-18 18-19 19-20

isolated ring systems :
 containing 1 : 15 : 25 :

G1:CH3,Et

G2:O,S

Match level :

1:Atom 2:Atom 3:Atom 4:Atom 5:Atom 6:Atom 7:CLASS 8:CLASS 10:CLASS 11:CLASS
 12:CLASS 14:CLASS 15:Atom 16:Atom 17:Atom 18:Atom 19:Atom 20:Atom 23:CLASS
 24:CLASS 25:Atom 26:Atom 27:Atom 28:Atom 29:Atom 30:CLASS 31:CLASS